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TRANSMITTAL LETTER TO THE UNITED STATES **DESIGNATED/ELECTED OFFICE (DO/EO/US) CONCERNING A SUBMISSION UNDER 35 U.S.C. 371**

ATTORNEY'S DOCKET NUMBER CS-8558/BCS033001

INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE PRIORITY DATE CLAIMED PCT/EP2004/000058 08 January 2004 (8.01.2004) TITLE OF INVENTION 9-KETOSPINOSYN DERIVATIVES 17 January 2003 (17.01.2003) APPLICANT(S) FOR DO/EO/US MALSAM, Olga; LOSEL, Peter; BECK, Michael E.; EBBINGHAUS-KINTSCHER, Ulrich; VELTEN, Robert; JESCHKE, Peter; MOHRLE, Volker; FRODE, Rita and EBERZ, Gunther Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: 1. X This is a FIRST submission of items concerning a submission under 35 U.S.C. 371. This is a SECOND or SUBSEQUENT submission of items concerning a submission under 35 U.S.C. 371. This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items The US has been elected (Article 31). A copy of the International Application as filed (35 U.S.C. 371(c)(2)) is attached hereto (required only if not communicated by the International Bureau). has been communicated by the International Bureau. is not required, as the application was filed in the United States Receiving Office (RO/US). \mathbf{X} An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)). is attached hereto. has been previously submitted under 35 U.S.C. 154(d)(4). Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3)) are attached hereto (required only if not communicated by the International Bureau). have been communicated by the International Bureau. have not been made; however, the time limit for making such amendments has NOT expired. have not been made and will not be made. An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)). 9. X An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)). An English language translation of the annexes of the International Preliminary Examination Report under PCT 10. Article 36 (35 U.S.C. 371(c)(5)). Items 11 to 20 below concern document(s) or information included: An Information Disclosure Statement under 37 CFR 1.97 and 1.98. 12. X An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included. A preliminary amendment. An Application Data Sheet under 37 CFR 1.76. A substitute specification. A power of attorney and/or change of address letter. A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821- 1.825. A second copy of the published International Application under 35 U.S.C. 154(d)(4). A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4). Preliminary Amendment w/Abstract Page; PTO Form 1449 and copies of foreign patents and articles listed therein Other items or information:

This collection of information is required by 37 CFR 1.414 and 1.491-1.492. The information is required to obtain or retain a benefit by the public, which is to file (and by the including gathering information, preparing, and submitting the completed form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount peparing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Commissioner for Patents P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Mail Stop PCT,

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Approved for use through 3/31/2007. OMB 0651-0021 U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number. U.S. APPLICATION NO. (if known, see 37 CFR 1.5) INTERNATIONAL APPLICATION NO. ATTORNEY'S DOCKET NUMBER PCT/EP2004/000058 CS-8558/BCS033001 CALCULATIONS PTO USE ONLY 21. 🔯 Basic national fee.....\$300 300.00 22. 🔀 If International preliminary examination report prepared by USPTO and all claims satisfy provisions of 200.00 All other situations \$200 23. X Search fee (37 CFR 1.445(a)(2)) has been paid on the international application to the USPTO as an International Searching Authority.....\$100 International Search Report prepared and provided to the Office......\$400 500.00 All other situations.....\$500 TOTAL OF 21, 22 and 23 = \$ Additional fee for specification and drawings filed in paper over 100 sheets (excluding 1,000.00 sequence listing or computer program listing filed in an electronic medium). The fee is \$250 for each additional 50 sheets of paper or fraction thereof. **Total Sheets** Extra Sheets Number of each additional 50 or fraction RATE thereof (round up to a whole number) 139 39 - 100 = /50 = 2 \$ x \$250 500.00 Surcharge of \$130.00 for furnishing the oath or declaration later than 30 months from the earliest claimed priority date (37 CFR 1.492(h)). \$ 0.00 **CLAIMS** NUMBER FILED NUMBER EXTRA RATE \$ Total claims - 20 = 14 × \$50 0 \$ 0.00 Independent claims 3 - 3 = 0 \$200 \$ 0.00 MULTIPLE DEPENDENT CLAIM(S) (if applicable) + \$360 \$ 0.00 TOTAL OF ABOVE CALCULATIONS = 1,500.00 Applicant claims small entity status. See 37 CFR 1.27. Fees above are reduced by 1/2. 0.00\$ 1,500.00 Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 CFR 1.492(i)). \$ 0.00 TOTAL NATIONAL FEE = \$ 1,500.00 Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property \$ 40.00 TOTAL FEES ENCLOSED = \$ 1,540.00 Amount to be \$ refunded: Amount to be \$ charged: A check in the amount of \$ to cover the above fees is enclosed. Please charge my Deposit Account No. 50-2510 in the amount of \$ 1,540.00 to cover the above fees. A duplicate copy of this sheet is enclosed. The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit c. [X] 50-2510 ... A duplicate copy of this sheet is enclosed. Fees are to be charged to a credit card. WARNING: Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038. NOTE: Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the International Application to pending status. SEND ALL CORRESPONDENCE TO: 34469 Richard E.L. Henderson NAME

31,619

REGISTRATION NUMBER

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TRANSMITTAL LETTER TO THE UNITED STATES RECEIVING OFFICES 41412

Express Mail mailing number: ED 780112807 NO. Deta of the city									
ED /80				1280	7 US	Date of deposit:	6 July 2005		
File reference no.: CS-8558/BCS 03-3001 International application no. (if known): PCT/EP2004/000058									
Title of the invention: 9-KETOSPINOSYN DERIVATIVES									
Earliest priority date claimed (Day/Month/Year): 17 Jan 2003 (17.01.2003)									
X This is a new International Application									
SCREENING DISCLOSURE INFORMATION:									
In order to assist in screening the accompanying international application for purposes of determining whether a license for foreign transmittal should and could be granted and for other purposes, the following information is supplied. (check as many boxes as apply):									
The in	distributed was not made in the Office States of America.								
There is no prior U.S. application relating to this invention.									
The fo	The following prior U.S. application(s) contain subject matter which is related to the invention disclosed in the attached international application. (NOTE: priority to these applications may or may not be claimed on the Request (form PCT/RO/101) and this listing does not constitute a claim for priority.)								
application					filed on			ו ר	
applicatio	n no.				filed on			1	
The present international application contains additional subject matter not found in the prior U.S. application(s) identified above. The additional subject matter is found on pages and DOES NOT ALTER MIGHT BE CONSIDERED TO ALTER the general nature of the invention in a manner which would require the U.S. application to have been made available for inspection by the appropriate defense agencies under 35 U.S.C. 181 and 37 C.F.R. 5.15. Itemized list of contents									
Sheets of Request form:					Charles				
Sheets of description					Check no.:				
(excluding sequence listing): 125				Return receipt postcard:					
Sheets of claims: 13				Power of attorney:					
Sheets of abstract:				Certified copy of priority document (specify):					
Sheets of drawings:				Other (specify): Preliminary Amendment w/Abstract Page;					
Sheets of sequence listing:			PTO Form 1449 and copies of foreign patents and articles						
Sequence listing diskette/CD:									
Tables related to sequence listing CD:									
The person signing this form is:		Applicant			Richard E.L. Henderson				
	X		ey/Agent (Reg. No.)		Name of person signing				
		31,619 Common Represen		Sin	enature Vila	1 2/2/1			

This collection of information is required by 37 CFR 1.10 and 1.412. The information is required to obtain or retain a benefit by the public, which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.11 and 1.14. This collection is estimated to take 15 minutes to complete, including gathering information, preparing, and submitting the completed form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Mail Stop PCT, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.